

Ref #	Hits	Search Query	DBs	Default Operator	Plurals	Time Stamp
L2	1	10/828343	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:41
L3	45	FLECKENSTEIN NEAR BERNHARD	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:37
L4	10	NEIPEL FRANK	US-PGPUB; USPAT; EPO; JPO; DERWENT	NEAR	ON	2006/01/26 14:37
L5	138	((HHV-8 or herpes\$10) WITH (interleukin-6 or IL-6 or V-IL-6 or vIL-6))	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:37
L6	10	((HHV-8 or herpes\$10) NEAR (interleukin-6 or IL-6 or V-IL-6 or vIL-6))	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:37
L7	7	((HHV-8 or herpes\$10) WITH (interleukin-6 or IL-6 or V-IL-6 or vIL-6)).clm.	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:37
L8	2	(V-IL-6 or vIL-6) NEAR (interleukin-6 or IL-6) NEAR receptor	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:38
L9	4	(V-IL-6 or vIL-6) SAME (interleukin-6 or IL-6) NEAR receptor	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:39
L10	335	Chang Yuan	US-PGPUB; USPAT; EPO; JPO; DERWENT	NEAR	OFF	2006/01/26 14:39
L11	7	L10 and L5	US-PGPUB; USPAT; EPO; JPO; DERWENT	NEAR	ON	2006/01/26 14:39
L12	20	Hayward Gary	US-PGPUB; USPAT; EPO; JPO; DERWENT	NEAR	OFF	2006/01/26 14:40
L13	1	L12 and L5	US-PGPUB; USPAT; EPO; JPO; DERWENT	NEAR	ON	2006/01/26 14:40
L14	2	GFNETSCLKKLADGFFEFE	US-PGPUB; USPAT; EPO; JPO; DERWENT	OR	ON	2006/01/26 14:41

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(FILE 'MEDLINE, CAPLUS, SCISEARCH, BIOSIS' ENTERED AT 14:48:30 ON 26 JAN 2006)

DEL HIS
E FLECKENSTEIN BERNHARD?/AU

L1 4 S E2
L2 230 S E1
L3 234 S L1 OR L2
L4 272 S V-IL-6 OR VIL-6
L5 159439 S INTERLEUKIN-6 OR IL-6
L6 211 S L4 (L) L5
L7 126 S L6 AND HHV?
L8 0 S L7 AND PY<=1996
L9 4 S L6 AND PY<=1996
L10 1 DUP REM L9 (3 DUPLICATES REMOVED)
L11 126 FOCUS L7 1-
L12 5 S L3 AND L11
L13 3 DUP REM L12 (2 DUPLICATES REMOVED)

=> d an ti so au ab pi 113 1-3

L13 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1
AN 1998:184856 CAPLUS
DN 128:293873
TI Human herpesvirus type 8 interleukin-6 homolog is functionally active on human myeloma cells
SO Blood (1998), 91(6), 1858-1863
CODEN: BLOOAW; ISSN: 0006-4971
AU Burger, Renate; Neipel, Frank; **Fleckenstein, Bernhard**; Savino, Rocco; Ciliberto, Gennaro; Kalden, Joachim R.; Gramatzki, Martin
AB Seroepidemiol. and polymerase chain reaction studies have strongly suggested that human herpesvirus type 8 (HHV-8) is associated with Kaposi's sarcoma, Castleman's disease, and body cavity-based lymphoma. The genome of HHV-8 harbors a viral analog of the interleukin-6 (IL-6) gene. The amino acid sequence of the viral IL-6 (vIL-6) protein is 24.7% identical to human IL-6 (hIL-6). IL-6 as a B-cell growth and differentiation factor is known to play an essential role in the pathophysiol. of B-cell tumors. Thus, it seems possible that virus-encoded IL-6 contributes to malignant growth of HHV-8-pos. B-cell lymphatic tumors. We have tested a preparation of HHV-8-derived IL-6 for the ability to promote the proliferation of the human myeloma cell line INA-6, which is strictly dependent on exogenous IL-6 for growth and survival. Viral IL-6 significantly induced DNA synthesis of INA-6 cells, but required much more protein on a weight basis when compared with hIL-6 for maximal proliferation. The proliferative effect of vIL-6 was almost completely inhibited by a combination of anti-IL-6 receptor (IL-6R) and anti-gp130 antibodies or IL-6R superantagonist Sant7 and anti-gp130 antibodies. This report demonstrates that vIL-6 has proliferative activity on human cells and that the IL-6R and gp130 are involved in vIL-6 signaling in the myeloma cell line INA-6.

L13 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
AN 1996:760040 CAPLUS
DN 126:55664
TI Human herpesvirus 8 encodes a homolog of interleukin-6
SO Journal of Virology (1997), 71(1), 839-842
CODEN: JOVIAM; ISSN: 0022-538X
AU Neipel, Frank; Albrecht, Jens-Christian; Ensser, Armin; Huang, Yao-Qi; Li, Jian Jun; Friedman-Kien, Alvin E.; **Fleckenstein, Bernhard**
AB Kaposi's sarcoma is a multifocal lesion that is reported to be greatly influenced by cytokines such as interleukin-6 (IL-6) and oncostatin M. DNA sequences of a novel human gammaherpesvirus, termed human herpesvirus 8 (HHV-8) or Kaposi sarcoma-associated herpesvirus, have been identified in all epidemiol. forms of Kaposi's sarcoma with high frequency. The presence of HHV-8

DNA is also clearly associated with certain B-cell lymphomas (body cavity-based lymphomas) and multicentric Castleman's disease. Sequence anal. of a 17-kb fragment revealed that adjacent to a block of conserved herpesvirus genes (major DNA-binding protein, glycoprotein B, and DNA polymerase), the genome of HHV-8 encodes structural homolog of IL-6. This cytokine is involved not only in the pathogenesis of Kaposi's sarcoma but also in certain B-cell lymphomas and multicentric Castleman's disease. The viral counterpart of IL-6 (vIL-6) has conserved important features such as cysteine residues involved in disulfide bridging or an amino-terminal signal peptide. Most notably, the region known to be involved in receptor binding is highly conserved in vIL-6. This conservation of essential features and the remarkable overlap between diseases associated with HIV-8 and diseases associated with IL-6 dysregulation clearly suggest that vIL-6 is involved in HHV-8 pathogenesis.

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S.Kaushal

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